#### AMENDMENT TO THE CLAIMS:

This listing of claims will replace all prior versions, and listings, of claims in the application:

#### Listing of Claims:

(Currently Amended) A method for inhibiting a mitotic kinesin Eg5 which comprises administering an effective amount of a thiadiazoline derivative represented by the general formula (I) or a pharmacologically acceptable salt thereof: A mitotic kinesin-Eg5 inhibitor which comprises a thiadiazoline derivative represented by the general formula (I) or a pharmacologically acceptable salt thereof as an active ingredient:

Output

Description:

√wherein:

R1 represents

- a hydrogen atom,
- a substituted or unsubstituted lower alkyl,
- a substituted or unsubstituted lower alkenyl,
- a substituted or unsubstituted lower alkynyl,
- a substituted or unsubstituted cycloalkyl.
- a substituted or unsubstituted aryl,
- or a substituted or unsubstituted heterocyclic group;

## R2 represents

- a hydrogen atom,
- a substituted or unsubstituted lower alkyl.
- a substituted or unsubstituted lower alkenyl,
- a substituted or unsubstituted lower alkynyl,
- a substituted or unsubstituted cycloalkyl,
- a substituted or unsubstituted aryl, or
- a substituted or unsubstituted heterocyclic group,
- -C(=W)R $^6$ , {wherein W represents an oxygen atom or a sulfur atom, and R $^6$  represents
  - a hydrogen atom.
  - a substituted or unsubstituted lower alkyl.
  - a substituted or unsubstituted lower alkenyl.
  - a substituted or unsubstituted lower alkynyl,
  - a substituted or unsubstituted cycloalkyl,
  - a substituted or unsubstituted aryl, or
  - a substituted or unsubstituted heterocyclic group,
  - -NR  $^7R^8_{\ \ \ \ }$  (wherein  $R^7$  and  $R^8$  are the same or different and
    - each represents a hydrogen atom,
    - a substituted or unsubstituted lower alkyl,
    - a substituted or unsubstituted lower alkenyl,
    - a substituted or unsubstituted lower alkynyl,
    - a substituted or unsubstituted cycloalkyl,

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\underline{a} substituted or unsubstituted aryl, or a substituted or unsubstituted heterocyclic group, or R^7 and R^8 are combined together with the adjacent nitrogen atom to form a substituted or unsubstituted heterocyclic group), -OR^9, \text{ (wherein } R^9 \text{ represents } \underline{a} \text{ substituted or unsubstituted lower alkyl,}
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- -OR<sup>9</sup>, (wherein R<sup>9</sup> represents <u>a</u> substituted or unsubstituted lower alkyl, <u>a</u> substituted or unsubstituted lower alkenyl, <u>a</u> substituted or unsubstituted lower alkynyl, <u>a</u> substituted or unsubstituted cycloalkyl, <u>a</u> substituted or unsubstituted aryl, or a substituted or unsubstituted heterocyclic group), or
- -SR  $^{10}$  , (wherein R  $^{10}$  has the same meaning as that of the aforementioned R  $^{9}$  H,
- -NR  $^{11}$ R  $^{12}$  , wherein R  $^{11}$  and R  $^{12}$  are the same or different and each represents a hydrogen atom,
  - a substituted or unsubstituted lower alkyl,
  - a substituted or unsubstituted lower alkenyl,
  - a substituted or unsubstituted lower alkynyl,
  - a substituted or unsubstituted cycloalkyl,
  - a substituted or unsubstituted aryl,
  - a substituted or unsubstituted heterocyclic group,
  - -C(=O)R13 {wherein R13 represents

- a hydrogen atom,
- a substituted or unsubstituted lower alkyl,
- a substituted or unsubstituted lower alkenyl,
- a substituted or unsubstituted lower alkynyl,
- a substituted or unsubstituted cycloalkyl,
- a substituted or unsubstituted aryl,
- a substituted or unsubstituted heterocyclic group,
- -NR<sup>14</sup>R<sup>15</sup><sub>a</sub> (wherein R<sup>14</sup> and R<sup>15</sup> are the same or different and each represents a hydrogen atom,
  - a substituted or unsubstituted lower alkyl,
  - a substituted or unsubstituted lower alkenyl,
  - a substituted or unsubstituted lower alkynyl,
  - a substituted or unsubstituted cycloalkyl,
  - a substituted or unsubstituted aryl, or
  - a substituted or unsubstituted heterocyclic group, or
  - R14 and R15 are combined together with the adjacent
    - nitrogen atom to form a substituted or unsubstituted heterocyclic group).
- -OR<sup>16</sup>, (wherein R<sup>16</sup> has the same meaning as that of the aforementioned R<sup>9</sup>), or
- -SR<sup>17</sup>, (wherein R<sup>17</sup> has the same meaning as that of the aforementioned R<sup>9</sup>), or

 $R^{11}$  and  $R^{12}$  are combined together with the adjacent nitrogen atom to form a substituted or unsubstituted heterocyclic group}, or

-SO<sub>2</sub>R<sup>18</sup>, (wherein R<sup>18</sup> represents

- a substituted or unsubstituted lower alkyl,
- a substituted or unsubstituted lower alkenyl,
- a substituted or unsubstituted lower alkynyl.
- a substituted or unsubstituted cycloalkyl.
- a substituted or unsubstituted aryl, or
- a substituted or unsubstituted heterocyclic group), or

 $R^1$  and  $R^2$  are combined together with the adjacent nitrogen atom to form a substituted or unsubstituted heterocyclic group,

## R3 represents

- a hydrogen atom, or
- -C(=Z)R  $^{19}_{-1}$  {wherein Z represents an oxygen atom or a sulfur atom, and R  $^{19}$  represents a hydrogen atom,
  - a substituted or unsubstituted lower alkyl,
  - a substituted or unsubstituted lower alkenyl,
  - a substituted or unsubstituted lower alkynyl,
  - $\underline{a}$  substituted or unsubstituted cycloalkyl,
  - a substituted or unsubstituted aryl,
  - a substituted or unsubstituted heterocyclic group, or
- -NR  $^{20}R^{21}$  , (wherein  $R^{20}$  and  $R^{21}$  are the same or different and each represents a hydrogen atom,

- a substituted or unsubstituted lower alkyl,
- a substituted or unsubstituted lower alkenyl.
- a substituted or unsubstituted lower alkynyl.
- a substituted or unsubstituted cycloalkyl.
- a substituted or unsubstituted arvl, or
- a substituted or unsubstituted heterocyclic group, or

R<sup>20</sup> and R<sup>21</sup> are combined together with the adjacent nitrogen atom to form a substituted or unsubstituted heterocyclic group),

# -OR<sup>22</sup> (wherein R<sup>22</sup> represents

- a substituted or unsubstituted lower alkyl,
- a substituted or unsubstituted lower alkenyl,
- a substituted or unsubstituted lower alkynyl,
- a substituted or unsubstituted cycloalkyl,
- a substituted or unsubstituted aryl, or
- a substituted or unsubstituted heterocyclic group), or
- -SR<sup>23</sup>, (wherein R<sup>23</sup> has the same meaning as that of the aforementioned R<sup>22</sup>)],

#### R4 represents

- a hydrogen atom,
- a substituted or unsubstituted lower alkyl,
- a substituted or unsubstituted lower alkenyl,
- a substituted or unsubstituted lower alkynyl,
- a substituted or unsubstituted cycloalkyl,
- a substituted or unsubstituted aryl, or

a substituted or unsubstituted heterocyclic group, and

# R5 represents

- a substituted or unsubstituted lower alkyl,
- $\underline{a}$  substituted or unsubstituted lower alkenyl,
- a substituted or unsubstituted lower alkynyl,
- a substituted or unsubstituted cycloalkyl,
- a substituted or unsubstituted aryl, or
- a substituted or unsubstituted heterocyclic group, or
- $R^4$  and  $R^5$  are combined together to represent -( $CR^{25A}R^{25B}$ )<sub>m1</sub>Q( $CR^{25C}R^{25D}$ )<sub>m2</sub>-, +wherein

Q represents a single bond, or

- a substituted or unsubstituted phenylene or cycloalkylene,
- m1 and m2 are the same or different and each represents an integer of from 0 to 4, with the proviso that m1 and m2 are not 0 at the same time.
- $R^{25A}$ ,  $R^{25B}$ ,  $R^{25C}$  and  $R^{25D}$  are the same or different and each represents a hydrogen atom,
  - a halogen,
  - a substituted or unsubstituted lower alkyl,
  - -OR26, wherein R26 represents
    - a hydrogen atom,
    - a substituted or unsubstituted lower alkyl.
    - a substituted or unsubstituted lower alkenyl,

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a substituted or unsubstituted lower alkynyl,

a substituted or unsubstituted cycloalkyl,

a substituted or unsubstituted aryl,

a substituted or unsubstituted heterocyclic group,

-CONR<sup>27</sup>R<sup>28</sup>, (wherein R<sup>27</sup> and R<sup>28</sup> are the same or

different and each represents

a hydrogen atom,

a substituted or unsubstituted lower alkyl,

<u>a</u> substituted or unsubstituted lower alkenyl,

<u>a</u> substituted or unsubstituted lower alkynyl,

a substituted or unsubstituted cycloalkyl,

a substituted or unsubstituted aryl, or

a substituted or unsubstituted heterocyclic group, or

 $R^{27}$  and  $R^{28}$  are combined together with the adjacent  $\label{eq:recombined} nitrogen \ atom \ to \ form \ a \ substituted \ or$ 

unsubstituted heterocyclic group),

 $-SO_2NR^{29}R^{30}$ , (wherein  $R^{29}$  and  $R^{30}$  have the same meanings as those of the aforementioned  $R^{27}$  and  $R^{28}$  respectively), or

-COR31, (wherein R31 represents

a hydrogen atom,

a substituted or unsubstituted lower alkyl,

a substituted or unsubstituted lower alkenyl,

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a substituted or unsubstituted lower alkynyl,
               a substituted or unsubstituted cycloalkyl,
               a substituted or unsubstituted arvl. or
               a substituted or unsubstituted heterocyclic group),
-NR32R33, twherein R32 and R33 are the same or different and each
     represents
       a hydrogen atom,
       a substituted or unsubstituted lower alkyl,
       a substituted or unsubstituted lower alkenyl,
       a substituted or unsubstituted lower alkynyl,
       a substituted or unsubstituted cycloalkyl,
       a substituted or unsubstituted aryl,
       a substituted or unsubstituted heterocyclic group,
       -COR34, (wherein R34 represents
               a hydrogen atom.
               a substituted or unsubstituted lower alkyl.
              a substituted or unsubstituted lower alkenyl.
              a substituted or unsubstituted lower alkynyl,
              a substituted or unsubstituted cycloalkyl,
               a substituted or unsubstituted aryl,
               a substituted or unsubstituted heterocyclic group,
              a substituted or unsubstituted lower alkoxy.
              a substituted or unsubstituted arvloxy, amino.
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a substituted or unsubstituted lower alkylamino.
                 a substituted or unsubstituted di-(lower alkyl)amino.
                 or a substituted or unsubstituted arylamino), or
        -SO<sub>2</sub>R<sup>35</sup>, (wherein R<sup>35</sup> represents
                 a substituted or unsubstituted lower alkyl.
                 a substituted or unsubstituted lower alkenyl.
                 a substituted or unsubstituted lower alkynyl.
                 a substituted or unsubstituted cycloalkyl.
                 a substituted or unsubstituted arvl, or
                a substituted or unsubstituted heterocyclic group).
or -COOR36, (wherein R36 represents
        a hydrogen atom,
        a substituted or unsubstituted lower alkyl,
        a substituted or unsubstituted lower alkenyl.
        a substituted or unsubstituted lower alkynyl.
        a substituted or unsubstituted cycloalkyl,
        a substituted or unsubstituted arvl, or
        a substituted or unsubstituted heterocyclic group), or
R<sup>25A</sup> and R<sup>25B</sup>, or R<sup>25C</sup> and R<sup>25D</sup> are combined together to represent
        an oxygen atom, and
when m1 or m2 is an integer of 2 or above, any of R25A, R25B, R25C
        and R25D may be the same or different, and any two of R25A.
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R<sup>25B</sup>, R<sup>25C</sup> and R<sup>25D</sup> which are bound to the adjacent two carbon atoms may be combined to form a bond}>.

- (Currently Amended) The method The mitotic kinesin Eg5 inhibitor according to claim 1, wherein R<sup>2</sup> is -C(=W)R<sup>6</sup><sub>2</sub> (wherein W and R<sup>6</sup> have the same meanings as those mentioned above, respectively).
- (Currently Amended) <u>The method The mitotic kinesin Eg5 inhibitor</u> according to claim 2, wherein R<sup>6</sup> is <u>a</u> substituted or unsubstituted lower alkyl.
- (Currently Amended) The method The mitotic kinesin Eg5 inhibitor according to claim 1, wherein R<sup>3</sup> is -C(=Z)R<sup>19</sup>, (wherein Z and R<sup>19</sup> have the same meanings as those mentioned above, respectively).
- (Currently Amended) <u>The method</u> <u>The mitotic kinesin Eg5 inhibitor according</u> to claim 4, wherein R<sup>19</sup> is <u>a</u> substituted or unsubstituted lower alkyl.
- (Currently Amended) <u>The method The mittetic kinesin Eg5 inhibitor</u> according to claim 1, wherein R<sup>5</sup> is <u>a</u> substituted or unsubstituted aryl, or a substituted or unsubstituted aromatic heterocyclic group.
- (Currently Amended) <u>The method</u> The mitotic kinesin Eg5 inhibitor according to claim 1, wherein R<sup>5</sup> is <u>a</u> substituted or unsubstituted aryl.

- 8. (Currently Amended) <u>The method</u> <u>The mitotic kinesin Eg5 inhibitor-according</u> to claim 1, wherein R<sup>4</sup> is <u>a</u> substituted or unsubstituted lower alkyl, or -(CH<sub>2</sub>)<sub>n</sub>NHSO<sub>2</sub>R<sup>24</sup>, (wherein n represents 1 or 2, and R<sup>24</sup> represents <u>a</u> substituted or unsubstituted lower alkyl, <u>a</u> substituted or unsubstituted lower alkenyl, <u>an</u> amino, <u>a</u> lower alkylamino, or <u>a</u> di-(lower alkyl)amino).
- 9. (Currently Amended) <u>The method</u> The mitotic-kinesin Eg5-inhibitor according to claim 1, wherein R<sup>4</sup> and R<sup>5</sup> are combined together to represent -(CR<sup>25A</sup>R<sup>25B</sup>)<sub>m1</sub>Q(CR<sup>25C</sup>R<sup>25D</sup>)<sub>m2\*</sub>, (wherein R<sup>25A</sup>, R<sup>25B</sup>, R<sup>25C</sup>, R<sup>25D</sup>, m1, m2, and Q have the same meanings as those mentioned above, respectively).
- (Currently Amended) <u>The method</u> <u>The mitotic kinesin Eg5-inhibitor</u>
   according to claim 9, wherein O is a substituted or unsubstituted phenylene.
- (Currently Amended) The method The mitotic kinesin Eg5 inhibitor according to claim 1, wherein R<sup>1</sup> is a hydrogen atom.
- (Currently Amended) <u>The method The mitotic kinesin Eg5 inhibitor</u>
   according to claim 1, wherein W and Z are oxygen atoms.
- (Currently Amended) A thiadiazoline derivative represented by the general formula (IA) or a pharmacologically acceptable salt thereof:

wherein R<sup>1A</sup> represents a hydrogen atom,

R2A represents

a hydrogen atom or

-COR<sup>6A</sup>, (wherein R<sup>6A</sup> represents <u>a</u> substituted or unsubstituted lower alkyl), or R<sup>1A</sup> and R<sup>2A</sup> are combined together with the adjacent nitrogen atom to form a substituted or unsubstituted heterocyclic group.

 $R^{3A}$  represents -COR<sup>19A</sup>, (wherein  $R^{19A}$  represents  $\underline{a}$  substituted or unsubstituted lower alkyl),

R<sup>4A</sup> represents

-(CH<sub>2</sub>)<sub>n</sub>NR<sup>4AA</sup>R<sup>4AB</sup>, fwherein

p represents 1 or 2, and

 $\boldsymbol{R}^{\text{4AA}}$  and  $\boldsymbol{R}^{\text{4AB}}$  are the same or different and each represents

a hydrogen atom,

a lower alkyl or cycloalkyl, (with the proviso that when R2A is -

 $\text{COR}^{6\text{A}}, \, \text{R}^{6\text{A}}$  and  $\text{R}^{19\text{A}}$  are tert-butyl and  $\text{R}^{5\text{A}}$  is phenyl,  $\text{R}^{4\text{AA}}$ 

and R<sup>4AB</sup> are not methyl at the same time),

-(CH2)pNR  $^{4AD}COR^{4AC}_{\ \ \ \ \ \ }$  (wherein p has the same meaning as that mentioned

above, R<sup>4AC</sup> represents a hydrogen atom, <u>a</u> lower alkyl, or <u>a</u> lower alkoxy,

and R<sup>4AD</sup> represents a hydrogen atom or a lower alkyl), or

-(CH<sub>2</sub>)<sub>p</sub>NHSO<sub>2</sub>R<sup>24A</sup>, (wherein p has the same meaning as that mentioned above,

R<sup>24A</sup> represents

-(CH<sub>2</sub>)<sub>q</sub>NR<sup>24AA</sup>, fwherein q represents an integer of from 0 to 5, and R<sup>24AA</sup> and R<sup>24AB</sup> are the same or different and each represents a hydrogen atom, <u>a</u> substituted or unsubstituted lower alkyl or cycloalkyl, (with the proviso that when R<sup>2A</sup> is -COR<sup>6A</sup>, R<sup>6A</sup> is tert-butyl and R<sup>19A</sup> is methyl or tert-butyl, neither of R<sup>24AA</sup> and R<sup>24AB</sup> is methyl, and if one of R<sup>24AA</sup> and R<sup>24AB</sup> is a hydrogen atom in-this-ease, the other is not ethyl or hydroxyethyl.

3-chloropropyl,

3-azidopropyl, or

lower alkenyl, (with the proviso that when  $R^{2A}$  is  $-COR^{6A}$ ,  $R^{6A}$  is tert-butyl and  $R^{19A}$  is methyl or tert-butyl,  $R^{24A}$  is not vinyl}, and

R<sup>5A</sup> represents <u>a</u> substituted or unsubstituted aryl or a substituted or unsubstituted aromatic heterocyclic group>.

- 14. (Original) The thiadiazoline derivative or a pharmacologically acceptable salt thereof according to claim 13, wherein R<sup>5A</sup> is substituted or unsubstituted aryl.
- 15. (Original) The thiadiazoline derivative or a pharmacologically acceptable salt thereof according to claim 13, wherein R<sup>5A</sup> is phenyl.

- 16. (Previously Presented) The thiadiazoline derivative or a pharmacologically acceptable salt thereof according to claim 13, wherein  $R^{2A}$  is  $COR^{6A}$ , and  $R^{6A}$  is unsubstituted lower alkyl.
- 17. (Previously Presented) The thiadiazoline derivative or a pharmacologically acceptable salt thereof according to claim 13, wherein R<sup>2A</sup> is COR<sup>6A</sup>, and R<sup>6A</sup> is tertbutyl.
- 18. (Previously Presented) The thiadiazoline derivative or a pharmacologically acceptable salt thereof according to claim 13, wherein R<sup>19A</sup> is unsubstituted lower alkyl.
- (Previously Presented) The thiadiazoline derivative or a pharmacologically acceptable salt thereof according to claim 13, wherein R<sup>19A</sup> is tert-butyl.
- 20. (Currently Amended) The thiadiazoline derivative or a pharmacologically acceptable salt thereof according to claim 13, wherein R<sup>4A</sup> is -(CH<sub>2</sub>)<sub>p</sub>NR<sup>4AA</sup>R<sup>4AB</sup> (wherein p, R<sup>4AA</sup> and R<sup>4AB</sup> have the same meanings as those mentioned above, respectively).
- 21. (Currently Amended) The thiadiazoline derivative or a pharmacologically acceptable salt thereof according to claim 13, wherein  $R^{4A}$  is -(CH<sub>2</sub>)<sub>p</sub>NR<sup>4AD</sup>COR<sup>4AC</sup><sub>a</sub> (wherein p,  $R^{4AC}$  and  $R^{4AD}$  have the same meanings as those mentioned above, respectively).

- 22. (Previously Presented) The thiadiazoline derivative or a pharmacologically acceptable salt thereof according to claim 13, wherein  $R^{4A}$  is  $-(CH_2)_pNHSO_2R^{24A}_2$  (wherein p and  $R^{24A}$  have the same meanings as those mentioned above, respectively).
- 23. (Previously Presented) A medicament which comprises the thiadiazoline derivative or a pharmacologically acceptable salt thereof according to claim 13 as an active ingredient.
  - 24. (Canceled)
  - 25. (Canceled)
- 26. (Previously Presented) A method for inhibiting a mitotic kinesin Eg5 which comprises administering an effective amount of the thiadiazoline derivative or a pharmacologically acceptable salt thereof according to claim 13.
  - 27.-28. (Canceled)